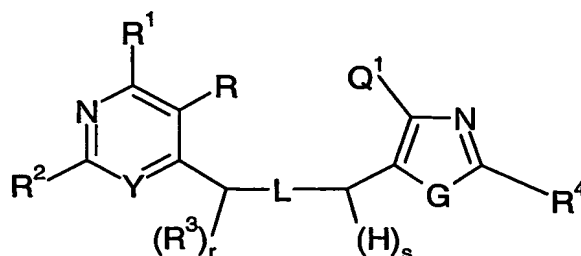


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CLAIMS

1. A compound of the Formula I:



Formula I

wherein:

**-L-** represents a double bond and r and s each represent 1 or **-L-** represents a triple bond and r and s each represent 0;

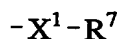
10 **G** is selected from O, S and NR<sup>5</sup>;

**Y** is selected from N and CR<sup>6</sup>;

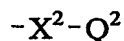
**Q<sup>1</sup>** is selected from aryl and heteroaryl,

and wherein **Q<sup>1</sup>** is optionally substituted by one or more substituents, which may be the same or different, selected from halogeno, trifluoromethyl, trifluoromethoxy, cyano,

15 nitro, hydroxy, amino, carboxy, carbamoyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulf-sulfinyl, (1-6C)alkylsulf-sulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, (3-6C)alkenoylamino, N-(1-6C)alkyl-(3-6C)alkenoylamino, (3-6C)alkynoylamino, N-(1-6C)alkyl-(3-6C)alkynoylamino, N-(1-6C)alkylsulf-sulfamoyl, N,N-di-[(1-6C)alkyl]sulf-sulfamoyl, (1-6C)alkanesulf-sulfonylamino, N-(1-6C)alkyl-(1-6C)alkanesulf-sulfonylamino, from a group of the formula:

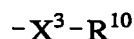


25 wherein X<sup>1</sup> is a direct bond or is selected from O and N(R<sup>8</sup>), wherein R<sup>8</sup> is hydrogen or (1-6C)alkyl, and R<sup>7</sup> is halogeno-(1-6C)alkyl, hydroxy-(1-6C)alkyl, (1-6C)alkoxy-(1-6C)alkyl, cyano-(1-6C)alkyl, amino-(1-6C)alkyl, (1-6C)alkylamino-(1-6C)alkyl or di-[(1-6C)alkyl]amino-(1-6C)alkyl, and from a group of the formula :



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wherein  $X^2$  is a direct bond or is selected from O, S, SO, SO<sub>2</sub>, N(R<sup>9</sup>), CO, CH(OR<sup>9</sup>), CON(R<sup>9</sup>), N(R<sup>9</sup>)CO, N(R<sup>9</sup>)CON(R<sup>9</sup>), SO<sub>2</sub>N(R<sup>9</sup>), N(R<sup>9</sup>)SO<sub>2</sub>, C(R<sup>9</sup>)<sub>2</sub>O, C(R<sup>9</sup>)<sub>2</sub>S and N(R<sup>9</sup>)C(R<sup>9</sup>)<sub>2</sub>, wherein R<sup>9</sup> is hydrogen or (1-6C)alkyl, and Q<sup>2</sup> is aryl, aryl-(1-6C)alkyl, heteroaryl, heteroaryl-(1-6C)alkyl, heterocyclyl or heterocyclyl-(1-6C)alkyl which optionally  
 5 bears 1, 2 or 3 substituents, which may be the same or different, selected from trifluoromethyl, trifluoromethoxy, cyano, nitro, hydroxy, amino, carboxy, carbamoyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, N-(1-6C)alkylsulfamoyl, N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino and N-(1-6C)alkyl-(1-6C)alkanesulfonylamino, or from a group of the formula:



- 15 wherein  $X^3$  is a direct bond or is selected from O and N(R<sup>11</sup>), wherein R<sup>11</sup> is hydrogen or (1-6C)alkyl, and R<sup>10</sup> is halogeno-(1-6C)alkyl, hydroxy-(1-6C)alkyl, (1-6C)alkoxy-(1-6C)alkyl, cyano-(1-6C)alkyl, amino-(1-6C)alkyl, (1-6C)alkylamino-(1-6C)alkyl or di-[(1-6C)alkyl]amino-(1-6C)alkyl, and any heterocyclyl group within Q<sup>2</sup> optionally bears 1 or 2 oxo or thioxo substituents;
- 20 **R** is selected from hydrogen, amino, hydroxy, halogeno, (1-6C)alkyl, (1-6C)alkoxy, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, carboxy, (1-6C)alkoxycarbonyl and *N*-(heterocyclyl(3-8C)cycloalkyl)carbamoyl;
- R<sup>1</sup>** is selected from hydrogen, halogeno, trifluoromethyl, trifluoromethoxy, cyano, nitro, hydroxy, amino, mercapto, carbamoyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, (3-6C)alkenoylamino, N-(1-6C)alkyl-(3-6C)alkenoylamino, (3-6C)alkynoylamino, N-(1-6C)alkyl-(3-6C)alkynoylamino, N-(1-6C)alkylsulfamoyl, N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino and N-(1-6C)alkyl-(1-6C)alkanesulfonylamino;
- 30

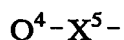
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$R^2$  is selected from hydrogen, halogeno, amino, hydroxy, halogeno, (1-6C)alkyl, (1-6C)alkoxy, (1-6C)alkylthio, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, aryl(1-6C)alkylamino, arylamino, heterocyclyl and (2-6C)alkanoylamino;

5  $R^3$  is selected from hydrogen, (1-6C)alkyl, hydroxy(1-6C)alkyl, carboxy, (1-6C)alkoxycarbonyl, carbamoyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl and *N*-(heterocyclyl(3-8C)cycloalkyl)carbamoyl;

10  $R^5$  is, independently, as defined for  $R^4$  and  $R^6$ , provided that  $R^5$  is not halogeno;

$R^4$  and  $R^6$  which may be the same or different, are selected from hydrogen, halogeno, trifluoromethyl, trifluoromethoxy, cyano, isocyano, nitro, hydroxy, mercapto, amino, formyl, carboxy, carbamoyl, sulfamoyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-15 6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, (3-6C)alkenoylamino, N-(1-6C)alkyl-(3-6C)alkenoylamino, (3-6C)alkynoylamino, N-(1-6C)alkyl-(3-6C)alkynoylamino, N-(1-20 6C)alkylsulfamoyl, N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino and N-(1-6C)alkyl-(1-6C)alkanesulfonylamino, or from a group of the formula :



wherein  $X^5$  is a direct bond or is selected from O, S, SO, SO<sub>2</sub>, N(R<sup>12</sup>), CO, CH(OR<sup>12</sup>), CON(R<sup>12</sup>), N(R<sup>12</sup>)CO, SO<sub>2</sub>N(R<sup>12</sup>), N(R<sup>12</sup>)SO<sub>2</sub>, OC(R<sup>12</sup>)<sub>2</sub>, SC(R<sup>12</sup>)<sub>2</sub> and N(R<sup>12</sup>)C(R<sup>12</sup>)<sub>2</sub>,

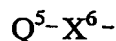
25 wherein R<sup>12</sup> is hydrogen or (1-6C)alkyl, and Q<sup>4</sup> is aryl, aryl-(1-6C)alkyl, (3-7C)cycloalkyl, (3-7C)cycloalkyl-(1-6C)alkyl, (3-7C)cycloalkenyl, (3-7C)cycloalkenyl-(1-6C)alkyl, heteroaryl, heteroaryl-(1-6C)alkyl, heterocyclyl or heterocyclyl-(1-6C)alkyl,

and wherein adjacent carbon atoms in any (2-6C)alkylene chain within an R<sup>4</sup>, R<sup>5</sup> or R<sup>6</sup> substituent are optionally separated by the insertion into the chain of a group selected from O, 30 S, SO, SO<sub>2</sub>, N(R<sup>13</sup>), CO, CH(OR<sup>13</sup>), CON(R<sup>13</sup>), N(R<sup>13</sup>)CO, SO<sub>2</sub>N(R<sup>13</sup>), N(R<sup>13</sup>)SO<sub>2</sub>, CH=CH and C≡C wherein R<sup>13</sup> is hydrogen or (1-6C)alkyl,

and wherein any CH<sub>2</sub>=CH- or HC≡C- group within an R<sup>4</sup>, R<sup>5</sup> or R<sup>6</sup> substituent optionally bears at the terminal CH<sub>2</sub>= or HC≡ position a substituent selected from halogeno,

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carboxy, carbamoyl, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, amino-(1-6C)alkyl, (1-6C)alkylamino-(1-6C)alkyl and di-[(1-6C)alkyl]amino-(1-6C)alkyl or from a group of the formula :



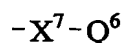
- 5 wherein  $X^6$  is a direct bond or is selected from CO and  $N(R^{14})CO$ , wherein  $R^{14}$  is hydrogen or (1-6C)alkyl, and  $Q^5$  is aryl, aryl-(1-6C)alkyl, heteroaryl, heteroaryl-(1-6C)alkyl, heterocyclyl or heterocyclyl-(1-6C)alkyl,

and wherein any  $CH_2$  or  $CH_3$  group within a  $R^4$ ,  $R^5$  or  $R^6$  substituent optionally bears on each said  $CH_2$  or  $CH_3$  group one or more halogeno or (1-6C)alkyl substituents or a

- 10 substituent selected from hydroxy, cyano, amino, carboxy, carbamoyl, (1-6C)alkoxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino,

(1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-

- 15 (2-6C)alkanoylamino, N-(1-6C)alkylsulfamoyl, N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino and N-(1-6C)alkyl-(1-6C)alkanesulfonylamino, or from a group of the formula :



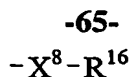
wherein  $X^7$  is a direct bond or is selected from O, S, SO,  $SO_2$ ,  $N(R^{15})$ , CO,  $CH(OR^{15})$ ,

- 20  $CON(R^{15})$ ,  $N(R^{15})CO$ ,  $SO_2N(R^{15})$ ,  $N(R^{15})SO_2$ ,  $C(R^{15})_2O$ ,  $C(R^{15})_2S$  and  $N(R^{15})C(R^{15})_2$ ,

wherein  $R^{15}$  is hydrogen or (1-6C)alkyl, and  $Q^6$  is aryl, aryl-(1-6C)alkyl, (3-7C)cycloalkyl, (3-7C)cycloalkyl-(1-6C)alkyl, (3-7C)cycloalkenyl, (3-7C)cycloalkenyl-(1-6C)alkyl, heteroaryl, heteroaryl-(1-6C)alkyl, heterocyclyl or heterocyclyl-(1-6C)alkyl,

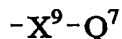
and wherein any aryl, heteroaryl, heterocyclyl, cycloalkyl or cycloalkenyl group

- 25 within a substituent on  $R^4$ ,  $R^5$  or  $R^6$  optionally bears 1 or more substituents, which may be the same or different, selected from halogeno, trifluoromethyl, trifluoromethoxy, cyano, nitro, hydroxy, amino, carboxy, carbamoyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, N-(1-6C)alkylsulfamoyl, N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino, N-(1-6C)alkyl-(1-6C)alkanesulfonylamino, from a group of the formula :



wherein  $X^8$  is a direct bond or is selected from O and  $N(R^{17})$ , wherein  $R^{17}$  is hydrogen or (1-6C)alkyl, and  $R^{16}$  is halogeno-(1-6C)alkyl, hydroxy-(1-6C)alkyl, (1-6C)alkoxy-(1-6C)alkyl, cyano-(1-6C)alkyl, amino-(1-6C)alkyl, (1-6C)alkylamino-(1-6C)alkyl, di-[(1-

5 6C)alkyl]amino-(1-6C)alkyl, (2-6C)alkanoylamino-(1-6C)alkyl or (1-6C)alkoxycarbonylamino-(1-6C)alkyl, and from a group of the formula :



wherein  $X^9$  is a direct bond or is selected from O, S, SO,  $SO_2$ ,  $N(R^{18})$ , CO,  $CH(OR^{18})$ ,  $CON(R^{18})$ ,  $N(R^{18})CO$ ,  $SO_2N(R^{18})$ ,  $N(R^{18})SO_2$ ,  $C(R^{18})_2O$ ,  $C(R^{18})_2S$  and  $N(R^{18})C(R^{18})_2$ ,

10 wherein  $R^{18}$  is hydrogen or (1-6C)alkyl, and  $Q^7$  is aryl, aryl-(1-6C)alkyl, (3-7C)cycloalkyl, (3-7C)cycloalkyl-(1-6C)alkyl, (3-7C)cycloalkenyl, (3-7C)cycloalkenyl-(1-6C)alkyl, heteroaryl, heteroaryl-(1-6C)alkyl, heterocyclyl or heterocyclyl-(1-6C)alkyl which optionally bears 1 or 2 substituents, which may be the same or different, selected from halogeno, trifluoromethyl, trifluoromethoxy, cyano, nitro, hydroxy, amino, carboxy, carbamoyl, (1-6C)alkyl, (2-15 8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, N-(1-6C)alkylsulfamoyl, N,N-di-[(1-6C)alkyl]sulfamoyl, (1-20 6C)alkanesulfonylamino and N-(1-6C)alkyl-(1-6C)alkanesulfonylamino,

or when G is  $NR^5$ ,  $R^4$  and  $R^5$  together with the atoms to which they are attached form a fused 5- or 6- membered heteroaryl or heterocyclyl ring, and wherein said fused 5- or 6-membered ring optionally bears one or more substituents as defined for  $R^4$ ,

25 and any fused 5- or 6- membered heterocyclyl ring so formed optionally bears 1 or 2 oxo or thioxo substituents,

and wherein any heterocyclyl group within any  $R^4$ ,  $R^5$  or  $R^6$  substituent optionally bears 1 or 2 oxo or thioxo substituents;  
or a pharmaceutically-acceptable salt thereof.

30

2. A pharmaceutical composition which comprises a compound of the Formula I, or a pharmaceutically acceptable salt thereof, as defined in claim 1 in association with a pharmaceutically-acceptable diluent or carrier.

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3. A compound of the Formula I, or a pharmaceutically-acceptable salt thereof, as defined in claim 1, for use as a medicament.

5 4. Use of a compound of the formula I, or a pharmaceutically acceptable salt thereof, as defined in claim 1, in the manufacture of a medicament for use as a Tie2 receptor tyrosine kinase inhibitor in a warm-blooded animal such as man.

5. Use of a compound of the formula I, or a pharmaceutically acceptable salt thereof, as  
10 defined in claim 1, in the manufacture of a medicament for use in the production of an anti-angiogenic effect in a warm-blooded animal such as man.